Amendments to the Claims

This listing of claims will replace all prior versions, and listing, of claims in the application.

1. **(Original)** A method for preparing a polyfunctionalized peptide comprising a peptidic backbone made up of four or more amino acids wherein two or more non-adjacent amino acids are independently substituted with a moiety having the structure:

with the proviso that the peptide sequence between any two consecutive, non-adjacent, amino acids bearing a A-L¹- moiety comprises at least one cysteine residue;

wherein the method comprises a step of:

reacting a peptide acyl donor comprising a peptidic backbone made up of two or more amino acids wherein said peptide acyl donor has the structure:

with a peptide amine acceptor having the structure:

$$R^{S1}S$$
 H_2N
 $Peptide Backbone$
 R^{X2}

under suitable conditions to effect ligation;

wherein k1 and k2 are independently integers between 1 and about 20;

each occurrence of A, A_1 and A_2 is independently an aliphatic, heteroaliphatic, aromatic, heteroaromatic, aryl, heteroaryl or a pharmaceutically useful group or entity;

R^{S1} is a sulfide protecting group;

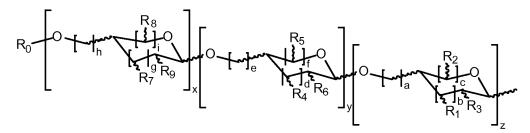
 R^{X0} is a group such that the moiety $-C(=O)OR^{X0}$ can be made to undergo ligation with the peptide amine acceptor;

each occurrence of L¹ is independently a substituted or unsubstituted, linear or branched, cyclic or acyclic, saturated or unsaturated aliphatic or heteroaliphatic moiety;

R^{X1} is hydrogen, alkyl, acyl, aromatic, heteroaromatic, aryl, heteroaryl, -alkyl(aryl), -alkyl(heteroaryl), a nitrogen protecting group, an amino acid or a proctected amino acid;

R^{X2} is -OR^{X2a} or -NR^{X2b}R^{X2c}, wherein R^{X2a} is hydrogen, alkyl, aromatic, heteroaromatic, aryl, heteroaryl, -alkyl(aryl), -alkyl(heteroaryl), a carboxylic acid protecting group, an amino acid or a proctected amino acid; and R^{X2b} and R^{X2c} are independently hydrogen, alkyl, aromatic, heteroaromatic, aryl, heteroaryl, -alkyl(aryl), -alkyl(heteroaryl), a nitrogen protecting group, an amino acid or a proctected amino acid.

- 2. **(Original)** The method of claim 1, wherein each occurrence of A, A1 and A2 is independently a pharmaceutically useful group or entity.
- 3. **(Original)** The method of claim 1, wherein each occurrence of A, A1 and A2 is independently a biomolecule, a small molecule, a macromolecule or a diagnostic label.
- 4. **(Original)** The method of claim 1, wherein each occurrence of A, A1 and A2 is independently a carbohydrate determinant having the structure:



wherein a, b, c, d, e, f, g, h, i, x, y and z are independently 0, 1, 2 or 3, with the proviso that the x, y and z bracketed structures represent furanose or pyranose moieties and the sum of b and c is 1 or 2, the sum of d and f is 1 or 2, and the sum of g and i is 1 or 2, and with the proviso that x, y and z are not simultaneously 0; wherein R_0 is hydrogen, a linear or branched chain alkyl, acyl, arylalkyl or aryl group; wherein each occurrence of R_1 , R_2 , R_3 , R_4 , R_5 , R_6 , R_7 , R_8 and

R₉ is independently hydrogen, OH, ORⁱ, NHRⁱ, NHCORⁱ, F, CH₂OH, CH₂ORⁱ, a substituted or unsubstituted linear or branched chain alkyl, (mono-, di- or tri)hydroxyalkyl, (mono-, di- or tri)acyloxyalkyl, arylalkyl or aryl group; wherein each occurrence of Rⁱ is independently hydrogen, CHO, COORⁱⁱ, or a substituted or unsubstituted linear or branched chain alkyl, acyl, arylalkyl or aryl group or a saccharide moiety having the structure:

wherein Y and Z are independently NH or O; wherein k, l, r, s, t, u, v and w are each independently 0, 1 or 2; with the proviso that the v and w bracketed structures represent furanose or pyranose moieties and the sum of l and k is 1 or 2, and the sum of s and u is 1 or 2, and with the proviso that v and w are not simultaneously 0; wherein R'₀ is hydrogen, a linear or branched chain alkyl, acyl, arylalkyl or aryl group; wherein each occurrence of R₁₀, R₁₁, R₁₂, R₁₃, R₁₄ and R₁₅ is independently hydrogen, OH, ORⁱⁱⁱ, NHRⁱⁱⁱ, NHCORⁱⁱⁱ, F, CH₂OH, CH₂ORⁱⁱⁱ, or a substituted or unsubstituted linear or branched chain alkyl, (mono-, di- or tri)hydroxyalkyl, (mono-, di- or tri)acyloxyalkyl, arylalkyl or aryl group; wherein each occurrence of R₁₆ is hydrogen, COOH, COORⁱⁱ, CONHRⁱⁱ, a substituted or unsubstituted linear or branched chain alkyl or aryl group; wherein each occurrence of Rⁱⁱⁱ is hydrogen, CHO, COOR^{iv}, or a substituted or unsubstituted linear or branched chain alkyl, acyl, arylalkyl or aryl group; and wherein each occurrence of Rⁱⁱⁱ and R^{iv} are each independently H, or a substituted or unsubstituted linear or branched chain alkyl, arylalkyl or aryl group.

- 5. **(Original)** The method of claim 1, wherein each occurrence of L^1 is independently $-O-(CH_2)_n$, wherein n is 0-9, or a glycoside-containing moiety.
- 6. (Original) The method of claim 1, wherein L^1 is -O-(CH_2)_n- CH_2 and two or more non-adjacent amino acids is/are independently substituted with a moiety having the structure:

Page 10 of 22

$$A$$
 C CH_2 S

wherein each occurrence of n is independently 0-8.

- 7. **(Original)** The method of claim 1, wherein each occurrence of A, A1 and A2 is independently selected from the group consisting of Globo-H, fucosyl GM1, KH-1, glycophorin, STN, (2,3)ST, Le^y, Le^x, N3, Tn, 2,6-STn, Gb3 and TF.
- 8. **(Original)** The method of claim 1, wherein the polyfunctionalized peptide has the structure:

$$\mathbb{R}^{X1} \left[\begin{array}{c} A_1 \\ A_2 \\ A_3 \\ A_4 \\ A_5 \\ A_7 \\ A_8 \\ A_9 \\ A_{13} \\ A_{14} \\ A_{15} \\ A_{15}$$

wherein s1 and s2 are independently an integer from 1 to about 20;

t1, t2 and t3 are each independently an integer;

R^{X1} is hydrogen, alkyl, acyl, aromatic, heteroaromatic, aryl, heteroaryl, -alkyl(aryl), -alkyl(heteroaryl), a nitrogen protecting group, an amino acid or a proctected amino acid;

R^{X2} is -OR^{X2a} or -NR^{X2b}R^{X2c}, wherein R^{X2a} is hydrogen, alkyl, aromatic, heteroaromatic, aryl, heteroaryl, -alkyl(aryl), -alkyl(heteroaryl), a carboxylic acid protecting group, an amino acid or a proctected amino acid; and R^{X2b} and R^{X2c} are independently hydrogen, alkyl, aromatic, heteroaromatic, aryl, heteroaryl, -alkyl(aryl), -alkyl(heteroaryl), a nitrogen protecting group, an amino acid or a proctected amino acid;

R^{P1}, R^{P2} and R^{P3} are independently H, alkyl, heteroalkyl, aromatic, heteroaromatic, aryl, heteroaryl, -alkyl(aryl), -alkyl(heteroaryl), or a natural or non-natural amino acid side chain;

each occurrence of L^1 is independently a substituted or unsubstituted aliphatic or heteroaliphatic moiety;

Page 11 of 22

 A_1 and A_2 are each independently an aliphatic, heteroaliphatic, aromatic, heteroaromatic, aryl, heteroaryl or a pharmaceutically useful group or entity; and

at least one occurrence of the bracketed structure t2 is a cysteine residue or protected cysteine residue;

and the method comprises a step of:

reacting a peptide acyl donor having the structure:

$$\begin{array}{c|c}
 & A_1 \\
\hline
 & A_1 \\
\hline
 & A_1
\end{array}$$

$$\begin{array}{c|c}
 & A_1 \\
\hline
 & A_1
\end{array}$$

$$\begin{array}{c|c}
 &$$

with a peptide amine acceptor having the structure:

under suitable conditions to effect ligation;

wherein the sum t+t' equals (t2)+1; R^{S1} is a sulfide protecting group; and R^{X0} is a group such that the moiety $-C(=O)OR^{X0}$ can be made to undergo ligation with the glycopeptide amine acceptor.

9. **(Original)** The method of claim 8, wherein the step of reacting the peptide acyl donor with the peptide amine acceptor is repeated a desired number of times, to prepare a polyfunctionalized peptide having the structure:

Page 12 of 22

$$\mathbb{R}^{X1} = \mathbb{R}^{P0} = \mathbb{R}^{P0} = \mathbb{R}^{P0} = \mathbb{R}^{P0} = \mathbb{R}^{P0} = \mathbb{R}^{P1} = \mathbb{R}$$

wherein R^{X1} and R^{X2} are as defined in claim 8;

each occurrence of A may be the same or different and may be as defined for A_1 and A_2 in claim 8;

each occurrence of R^{P1} may be the same or different and may be as defined for R^{P1} and R^{P2} in claim 8;

q is an integer greater than or equal to 2;

each occurrence of s is independently an integer from 1 to about 20;

each occurrence of t is independently an integer;

t0 is an integer; and

each occurrence of R^{P0} is independently H, alkyl, heteroalkyl, aromatic, heteroaromatic, aryl, heteroaryl, -alkyl(aryl), -alkyl(heteroaryl), or a natural or non-natural amino acid side chain.

- 10. (Original) The method of claim 9, wherein q is an integer between 2 and about 5.
- 11. **(Original)** The method of claim 9, wherein q is 2.
- 12. **(Original)** The method of claim 9, wherein the sum s+t is between about 2 and about 6.
- 13. (Original) The method of claim 9, wherein t0 is an integer from 0 to about 20.
- 14. (Original) The method of claim 9, wherein R^{X1} is hydrogen, Fmoc or Ac.
- 15. (Original) The method of claim 9, wherein R^{X2} is NH_2 .

- 16. (Original) The method of claim 9, wherein R^{X0} is disulfide-substituted aryl moiety.
- 17. (Original) The method of claim 9, wherein R^{X0} has the structure:

wherein R is an aliphatic, heteroaliphatic, aromatic or heteroaromatic moiety.

18. (Original) The method of claim 17, wherein R^{X0} has the structure:

wherein R is lower alkyl.

- 19. **(Original)** The method of claim 18, wherein R is ethyl.
- 20. (Original) The method of claim 9, wherein R^{S1} is -StBu.
- 21. **(Original)** The method of claim 9, wherein in the step of reacting the peptide acyl donor having the structure:

with the peptide amine acceptor under suitable conditions to effect ligation, an intermediate having the following structure is formed in situ:

$$\mathbb{R}^{X1} \begin{bmatrix} H & 0 \\ N & 1 \\ R^{P1} \end{bmatrix}_{t1} \begin{bmatrix} N & 0 \\ N & 1 \\ H & 0 \end{bmatrix}_{s1} \begin{bmatrix} H & 0 \\ N & 1 \\ R^{P2} \end{bmatrix}_{t} \mathbb{S}^{X0z}$$

wherein R^{X0a} is an oxygen-substituted aryl moiety.

- 22. **(Original)** The method of claim 21, wherein the suitable conditions to effect ligation comprise MESNa.
- 23. **(Original)** The method of claim 9, wherein in the peptide acyl donor having the structure:

$$\mathbb{R}^{X1} \begin{bmatrix} H & O \\ N & Y \\ R^{P1} \end{bmatrix}_{t1} \begin{bmatrix} N & A_1 \\ N & Y \\ H & O \end{bmatrix}_{s1} \begin{bmatrix} H & O \\ N & Y \\ R^{P2} \end{bmatrix}_{t} O\mathbb{R}^{X0}$$

the amino acyl residue directly attached to $-OR^{X0}$ is phenylalanine.

- 24. **(Original)** The method of claim 1, wherein when at least one occurrence of A (or A_1 and/or A_2 , as further defined for A) is a carbohydrate domain, some or all of carbohydrate domains are O-linked to the peptide backbone.
- 25. (Original) The method of claim 1, wherein when at least one occurrence of A (or A_1 and/or A_2 , as further defined for A) is a carbohydrate domain, some or all of carbohydrate domains are N-linked to the peptide backbone.

Page 15 of 22

- 26. **(Original)** The method of claim 1, wherein the polyfunctionalized peptide is symmetrical.
- 27. **(Original)** The method of claim 1, wherein the polyfunctionalized peptide is nonsymmetrical.
- 28. **(Original)** The method of claim 1, further comprising a step of conjugating the polyfunctionalized peptide to an immunogenic carrier.
- 29. (Original) The method of claim 28, wherein the carrier is a protein, a peptide or a lipid.
- 30. **(Original)** The method of claim 28, wherein the carrier is Bovine Serum Albumin (BSA), Keyhole Limpet Hemocyanin (KLH) or polylysine.
- 31. **(Original)** The method of claim 28, wherein the carrier is a lipid carrier having the structure:

wherein m, n and p are each independently integers between about 8 and 20; and $R_{\rm V}$ is hydrogen, substituted or unsubstituted linear or branched chain lower alkyl or substituted or unsubstituted phenyl.

- 32. **(Original)** The method of claim 31, wherein m', n' and p' are each 14.
- 33. **(Original)** The method of claim 28, wherein the carrier is linked to the polyfunctionalized peptide through a crosslinker.

Page 16 of 22

34. **(Original)** The method of claim 33, wherein the crosslinker is a fragment having the structure:

whereby said structure is generated upon conjugation of a maleimidobenzoic acid N-hydroxy succinimide ester with a suitable functionality on the polyfunctionalized peptide.

35. **(Currently Amended)** The method of claim 1, wherein the polyfunctionalized peptide has the structure:

36. **(Currently Amended)** The method of claim 1, wherein the polyfunctionalized peptide has the structure:

Page 17 of 22

37. **(Currently Amended)** The method of claim 1, wherein the polyfunctionalized peptide has the structure:

38. **(Currently Amended)** The method of claim 1, wherein the polyfunctionalized peptide has the structure:

39. **(Currently Amended)** The method of claim 1, wherein the polyfunctionalized peptide has the structure:

40. (Cancelled)